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Date: February 1, 2006

Time Sent:

To:	Company:	Facsimile No:	Telephone No:
Examiner Marianne Allen	U.S. Patent and Trademark Office	571.273.0712	571.272.0712
From:	Cameron K. Weiffenbach	Direct Phone:	202.756.8171
E-Mail:	cweiffenbach@mwe.com		
Sent By:	Jackie Reid-Johnson	Direct Phone:	202.756.8668
Client/Matter/Tkpr:	050179-0081/05169	Original to Follow by Mail:	No
		Number of Pages, Including Cover:	32

Message:

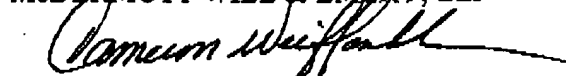
U.S. Patent Application No.09/555,275
Art Unit 1631
Applicant: Bentley et al.

Dear Examiner Allen:

Attached is a copy of a document filed in the USPTO on February 5, 2002 along with evidence of receipt of the document by the USPTO. The document is a substitute sequence listing and a preliminary amendment. This document should take care of the printer query. Should you have any questions, please do not hesitate to call me.

Sincerely yours,

McDERMOTT WILL & EMERY, LLP



Cameron K. Weiffenbach
Registration No. 44,488

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U.S. practice conducted through McDermott Will & Emery LLP.
600 Thirteenth Street, N.W. Washington, D.C. 20005-3096

Telephone: 202.756.8000

Dram Davis

Country Application

05-Feb-02

Client-Matter: 050179-0081 **Country:** US **SubCase:**
Family Number: 050179-0081 **United States of America**
Case Type: PCT **Application Status:** Pending
Application Number: 09/555,275 **Filing Date:** 26-May-2000
Patent Number: **Issue Date:**
Publication Number: **Publication Date:**
Priority Number: PP0585 **Priority Date:** 27-Nov-1997
Tax Schedule: LE **Expiration Date:**
Reel & Frame: 010994/0301 **Tax Start Date:**
Group Art Unit: **ClientRef:** 92546
Agent:
Agent Reference Number:

List Of User Actions

Action(s) Due	Due Date	Action Taken
Office Action Received Yet?	26-Feb-2002	Due Date
FILING RECEIPT RECD YET?	07-Mar-2002	Due Date

Applicant: John David BENTLEY, et al. Docket No. 50179-081
 Title/Mark: METHOD OF DESIGNING AGONISTS AND ANTAGONISTS TO
IGF RECEPTOR Serial/Reg./Patent No. 09/555,275
 Date Sent: 2/5/02 ☒ Hand Carried ☐ Fax ☐ Electronic ☐ Cert. of Mailing ☐ Express Mail No. _____
☐ Transmittal Letter
 New Patent App ☒ Utility ☐ Design ☐ Cont. ☐ CIP ☐ Div. ☐ PCT ☐ CPA ☐ RCE ☐ Prov
☐ Other: _____
 _____ pages of Specification
 _____ pages of Claims
 _____ pages of Abstract
 _____ pages of Formal/Informal Drawings
☐ Small Entity ☐ Large Entity
☐ Declaration/Power of Attorney
☐ Recordation of Assignment/Security Agreement
☐ Information Disclosure Statement
 Form PTO 1449
 _____ copies of cited references
☒ Preliminary Amendment
☐ Response to Missing Parts Notice
☐ Resp. to Notice to Correct App. Papers
☐ Certified Copy of Priority Doc.
☐ Claim for Convention Priority
☐ Response/Amendment to Office Action of _____
☐ Request for _____ day/month Extension of Time

☐ Letter submitting _____ pages of drawings
☐ Req. for Approval of Drawing Amendments
☐ Req. for Oral Hearing
☐ Not. of Appeal ☐ Appeal Brief ☐ Reply Brief
☐ Rule 312 Amendment/Letter
☐ Req. for Acknowledgement of Cited Art
☐ Issue Fee
☐ Publication Fee
☐ Req. for Certificate of Correction
☐ Maintenance Fee for _____ years after grant
☐ Fee Address Indication Form _____
☐ Terminal Disclaimer
☐ Petition to Commissioner
☐ Status Inquiry
☐ Other Submission of Substitute Sequence Listing, Preliminary Amendment, Diskette Containing Computer Readable
☒ Copy of Sequence Listing

O I P E
 FEB 05 2002
 PATENT & TRADEMARK OFFICE

ANTI-STATIC
MEDIA MAILER

Applicant: COMMONWEALTH SCIENTIFIC
AND INDUSTRIAL RESEARCH
ORGANISATION

Title: METHOD OF DESIGNING
AGONISTS AND ANTAGONISTS TO IGF
RECEPTOR

Attorney Docket: 050179-0081
Data Recorded: February 5, 2002

U.S. Serial NO.: 09/555,275
U.S. Filing Date: 26 May 2000

MS-DOS, ASCII Format

CAUTION

Do not bend or fold
Avoid exposure to all magnetic fields

Attorney Docket No. 050179-0081**PATENT APPLICATION****IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

In re the Application of)
)
John David BENTLEY, et al.)
) Group Art Unit: TBA
Serial No.: 09/555,275)
) Examiner: TBA
Filed: May 26, 2000)
)
For: METHOD OF DESIGNING AGONISTS AND)
ANTAGONISTS TO IGF RECEPTOR)

**SUBMISSION OF SUBSTITUTE SEQUENCE LISTING
AND PRELIMINARY AMENDMENT**

Honorable Assistant Commissioner for Patents
BOX SEQUENCE
Washington, D.C. 20231

Sir:

Prior to initial examination of the above-captioned patent application, please amend the above-captioned patent application as follows:

IN THE SPECIFICATION:

Please substitute the second paragraph on page 20 and continuing on to page 21 of the specification with the following rewritten paragraph.

-- Soluble IGF-1R/462 protein was recovered from harvested fermentation medium by affinity chromatography on columns prepared by coupling Mab 9E10 to divinyl sulphone-activated agarose beads (Mini Leak: Kem En Tec. Denmark) as recommended by the manufacturer. Mini-Leak Low and Medium affinity columns with antibody loadings of 1.5-4.5

WDC99 555265-1.050179.0081

Attorney Docket No. 050179-0081
Application Serial No. 09/555.275

mg/ml of hydrated matrix were obtained, with the loading range of 2.5-3 mg/ml giving optimal performance (data not shown). Mab 9E10 was produced by growing hybridoma cells (American Tissue Culture Collection) in serum-free medium in the Celligen Plus bioreactor and recovering the secreted antibody (4 g) using protein A glass beads (Prosep-A, bioprocessing Limited, USA). Harvested culture medium containing IGF-1R/462 protein was adjusted to pH 8.0 with Tris-HCl (Sigma), made 0.02% (w/v) in sodium azide and passed at 3-5 ml/min over 50 ml Mab 9E10 antibody columns at 4° C. Bound protein was recovered by recycling a solution of 2-10 mg of the undecamer c-myc peptide EQKLISEEDLN (SEQ ID NO. 16) (Hoogenboom et al., 1991) in 20 ml of Tris-buffered saline containing 0.02% sodium azide (TBSA). Between 65% and 75% of the product was recovered from the medium as estimated by ELISA, with a further 15-25% being recovered by a second pass over the columns. Peptide recirculation (~10 times) through the column eluted bound protein more efficiently than a single, slower elution. Residual bound protein was eluted with sodium citrate buffer at pH 3.0 into 1 M Tris HCl pH 8.0 to neutralize the eluant, and columns were re-equilibrated with TBSA.--

**Please insert after page 46 and before the claims the attached paper copy of the this
Substitute Sequence Listing.**

REMARKS

The specification is corrected and a Substitute Sequence Listing is herein submitted to

Attorney Docket No. 050179-0081
Application Serial No. 09/555,275

comply with the requirements for an application containing a nucleotide and/or amino acid sequence.

Hereto is an attached Substitute Sequence Listing in paper and computer readable format.
The paper copy and computer readable copy of the Substitute Sequence Listing are the same.
The substitute Sequence Listing does not include new matter.

CONCLUSION

Entry of the Substitute Sequence Listing and favorable consideration are respectfully requested.

To the extent necessary, please grant any extension of time deemed necessary for entry of this communication. Please charge any deficient fees, or credit any overpayment of fees, to Deposit Account 500417.

Respectfully submitted,

McDermott, Will & Emery



Kelli N. Watson

Registration No. 47,170

DATE: February 5, 2002

Attorney Docket No. 050179-0081
Application Serial No. 09/555,275

McDermott, Will & Emery
600 Thirteenth Street, N.W.
Washington, D.C. 20005-3096
(202) 756-8351 (Telephone, direct)
(202) 756-8087 (Facsimile)

Attachments:

Paper Copy of Sequence Listing
Diskette Containing Computer Readable
Copy of Sequence Listing

Attorney Docket No. 050179-0081
Application Serial No. 09/555,275

ATTACHMENT

Version With Markings To Show Changes Made

IN THE SPECIFICATION

The second paragraph on page 20 and continuing on to page 21 of the specification is substituted with the following rewritten paragraph.

— Soluble IGF-1R/462 protein was recovered from harvested fermentation medium by affinity chromatography on columns prepared by coupling Mab 9E10 to divinyl sulphone-activated agarose beads (Mini Leak: Kem En Tec. Denmark) as recommended by the manufacturer. Mini-Leak Low and Medium affinity columns with antibody loadings of 1.5-4.5 mg/ml of hydrated matrix were obtained, with the loading range of 2.5-3 mg/ml giving optimal performance (data not shown). Mab 9E10 was produced by growing hybridoma cells (American Tissue Culture Collection) in serum-free medium in the Celligen Plus bioreactor and recovering the secreted antibody (4 g) using protein A glass beads (Prosep-A, bioprocessing Limited, USA). Harvested culture medium containing IGF-1R/462 protein was adjusted to pH 8.0 with Tris-HCl (Sigma), made 0.02% (w/v) in sodium azide and passed at 3-5 ml/min over 50 ml Mab 9E10 antibody columns at 4° C. Bound protein was recovered by recycling a solution of 2-10 mg of the undecamer c-myc peptide EQKLISEEDLN (SEQ ID NO. 16) (Hoogenboom et al., 1991) in 20 ml of Tris-buffered saline containing 0.02% sodium azide (TBSA). Between 65% and 75% of the product was recovered from the medium as estimated by ELISA, with a further 15-25% being

Attorney Docket No. 050179-0081
Application Serial No. 09/555,275

recovered by a second pass over the columns. Peptide recirculation (~10 times) through the column eluted bound protein more efficiently than a single, slower elution. Residual bound protein was eluted with sodium citrate buffer at pH 3.0 into 1 M Tris HCl pH 8.0 to neutralize the eluant, and columns were re-equilibrated with TBSA.--

The attached paper copy of this Substitute Sequence Listing is inserted after page 46 and before the claims of the specification.

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<140> 09/555,275
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<150> PCT/AU98/00998
<151> 1998-11-27
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35 40 45

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50 55 60

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Leu Phe Tyr Asn Tyr Ala Leu Val Ile Phe Glu Met Thr Asn Leu Lys
85 90 95

Asp Ile Gly Leu Tyr Asn Leu Arg Asn Ile Thr Arg Gly Ala Ile Arg
100 105 110

Ile Glu Lys Asn Ala Asp Leu Cys Tyr Leu Ser Thr Val Asp Trp Ser
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Leu Ser Phe Pro Lys Leu Ile Met Ile Thr Asp Tyr Leu Leu Leu Phe
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Arg Val Tyr Gly Leu Glu Ser Leu Lys Asp Leu Phe Pro Asn Leu Thr
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 100 105 110

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Asn Tyr Asp Leu Ser Phe Leu Lys Thr Ile Gln Glu Val Ala Gly Tyr
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Val Leu Ile Ala Leu Asn Thr Val Glu Arg Ile Pro Leu Glu Asn Leu
65 70 75 80

Gln Ile Ile Arg Gly Asn Met Tyr Tyr Glu Asn Ser Tyr Ala Leu Ala
85 90 95

Val Leu Ser Asn Tyr Asp Ala Asn Lys Thr Gly Leu Xaa Xaa Lys Pro
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Met Arg Asn Leu Gln Glu Ile Leu His Gly Ala Val Arg Phe Ser Asn
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Lys Glu Ile Thr Gly Phe Leu Leu Ile Gln Ala Trp Pro Glu Asn Arg
65 70 75 80

Thr Asp Leu His Ala Phe Glu Asn Leu Glu Ile Ile Arg Gly Arg Thr
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Lys Gln His Gly Gln Phe Ser Leu Ala Val Val Ser Leu Asn Ile Thr
100 105 110

Ser Leu Gly Leu Arg Ser Leu Lys Glu Ile Ser Asp Gly Asp Val Ile
115 120 125

Ile Ser Gly Asn Lys Asn Leu Cys Tyr Ala Asn Thr Ile Asn Trp Lys
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50 55 60

Tyr Ala Leu Val Ser Leu Ser Phe Phe Arg Lys Leu Arg Leu Ile Arg
65 70 75 80

Gly Glu Thr Leu Glu Ile Gly Asn Tyr Ser Phe Tyr Ala Leu Asp Asn
85 90 95

Gln Asn Leu Arg Gln Leu Trp Asp Trp Ser Lys His Asn Leu Thr Ile
100 105 110

Thr Gln Gly Lys Leu Phe Phe His Tyr Asn Pro Lys Leu Cys Leu Ser
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 35 40 45

Met Gly Leu Ile Glu Val Val Thr Gly Tyr Val Lys Ile Arg His Ser
 50 55 60

His Ala Leu Val Ser Leu Ser Phe Leu Lys Asn Leu Arg Leu Ile Leu
 65 70 75 80

Gly Glu Glu Gln Leu Glu Gly Asn Tyr Ser Phe Tyr Val Leu Asp Asn
 85 90 95

Gln Asn Leu Gln Gln Leu Trp Asp Trp Asp His Arg Asn Leu Thr Ile
 100 105 110

Lys Ala Gly Lys Met Tyr Phe Ala Phe Asn Pro Lys Leu Cys Val Ser
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50 55 60

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Cys Val Glu Thr Cys Pro Pro Pro Tyr Tyr His Phe Gln Asp Trp Arg
85 90 95

Cys Val Asn Phe Ser Phe Cys Gln Asp Leu His His Lys Cys Lys Asn
100 105 110

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65 70 75 80

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755 760 765

Tyr Arg Ile Asp Ile His Ser Cys Asn His Glu Ala Glu Lys Leu Gly
770 775 780

Cys Ser Ala Ser Asn Phe Val Phe Ala Arg Thr Met Pro Ala Glu Gly
785 790 795 800

Ala Asp Asp Ile Pro Gly Pro Val Thr Trp Glu Pro Arg Pro Glu Asn
805 810 815

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835 840 845

Cys Val Ser Arg Gln Glu Tyr Arg Lys Tyr Gly Gly Ala Lys Leu Asn
850 855 860

Arg Leu Asn Pro Gly Asn Tyr Thr Ala Arg Ile Gln Ala Thr Ser Leu
865 870 875 880

Ser Gly Asn Gly Ser Trp Thr Asp Pro Val Phe Phe Tyr Val Gln Ala
885 890 895

Lys Thr Gly Tyr Glu Asn Phe Ile His Leu
900 905

<210> 12
<211> 916
<212> PRT
<213> Homo sapiens

<400> 12

His Leu Tyr Pro Gly Glu Val Cys Pro Gly Met Asp Ile Arg Asn Asn
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Leu Thr Arg Leu His Glu Leu Glu Asn Cys Ser Val Ile Glu Gly His
20 25 30

Leu Gln Ile Leu Leu Met Phe Lys Thr Arg Pro Glu Asp Phe Arg Asp
35 40 45

Leu Ser Phe Pro Lys Leu Ile Met Ile Thr Asp Tyr Leu Leu Leu Phe
50 55 60

Arg Val Tyr Gly Leu Glu Ser Leu Lys Asp Leu Phe Pro Asn Leu Thr
65 70 75 80

Val Ile Arg Gly Ser Arg Leu Phe Phe Asn Tyr Ala Leu Val Ile Phe
85 90 95

Glu Met Val His Leu Lys Glu Leu Gly Leu Tyr Asn Leu Met Asn Ile
100 105 110

Thr Arg Gly Ser Val Arg Ile Glu Lys Asn Asn Glu Leu Cys Tyr Leu
115 120 125

Ala Thr Ile Asp Trp Ser Arg Ile Leu Asp Ser Val Glu Asp Asn Tyr
130 135 140

Ile Val Leu Asn Asp Asp Asn Glu Glu Cys Gly Asp Ile Cys Pro Gly
145 150 155 160

Thr Ala Lys Gly Lys Thr Asn Cys Pro Ala Thr Val Ile Asn Gly Gln
165 170 175

Phe Val Glu Arg Cys Trp Thr His Ser His Cys Gln Lys Val Cys Pro
180 185 190

Thr Ile Cys Lys Ser His Gly Cys Thr Ala Glu Gly Leu Cys Cys His
195 200 205

Ser Glu Cys Leu Gly Asn Cys Ser Gln Pro Asp Asp Pro Thr Lys Cys
210 215 220

Val Ala Cys Arg Asn Phe Tyr Leu Asp Gly Arg Cys Val Glu Thr Cys
225 230 235 240

Pro Pro Pro Tyr Tyr His Phe Gln Asp Trp Arg Cys Val Asn Phe Ser
245 250 255

Phe Cys Gln Asp Leu His His Lys Cys Lys Asn Ser Arg Arg Gln Gly
260 265 270

Cys His Gln Tyr Val Ile His Asn Asn Lys Cys Ile Pro Glu Cys Pro
275 280 285

Ser Gly Tyr Thr Met Asn Ser Ser Asn Leu Leu Cys Thr Pro Cys Leu
290 295 300

Gly Pro Cys Pro Lys Val Cys His Leu Leu Glu Gly Glu Lys Thr Ile
305 310 315 320

Asp Ser Val Thr Ser Ala Gln Glu Leu Arg Gly Cys Thr Val Ile Asn
325 330 335

Gly Ser Leu Ile Ile Asn Ile Arg Gly Gly Asn Asn Leu Ala Ala Glu
340 345 350

Leu Glu Ala Asn Leu Gly Leu Ile Glu Glu Ile Ser Gly Tyr Leu Lys
355 360 365

Ile Arg Arg Ser Tyr Ala Leu Val Ser Leu Ser Phe Phe Arg Lys Leu
370 375 380

Arg Leu Ile Arg Gly Glu Thr Leu Glu Ile Gly Asn Tyr Ser Phe Tyr
385 390 395 400

Ala Leu Asp Asn Gln Asn Leu Arg Gln Leu Trp Asp Trp Ser Lys His
405 410 415

Asn Leu Thr Ile Thr Gln Gly Lys Leu Phe Phe His Tyr Asn Pro Lys
420 425 430

Leu Cys Leu Ser Glu Ile His Lys Met Glu Glu Val Ser Gly Thr Lys
435 440 445

Gly Arg Gln Glu Arg Asn Asp Ile Ala Leu Lys Thr Asn Gly Asp Gln
450 455 460

Ala Ser Cys Glu Asn Glu Leu Leu Lys Phe Ser Tyr Ile Arg Thr Ser
465 470 475 480

Phe Asp Lys Ile Leu Leu Arg Trp Glu Pro Tyr Trp Pro Pro Asp Phe
485 490 495

Arg Asp Leu Leu Gly Phe Met Leu Phe Tyr Lys Glu Ala Pro Tyr Gln
500 505 510

Asn Val Thr Glu Phe Asp Gly Gln Asp Ala Cys Gly Ser Asn Ser Trp
515 520 525

Thr Val Val Asp Ile Asp Pro Pro Leu Arg Ser Asn Asp Pro Lys Ser
530 535 540

Gln Asn His Pro Gly Trp Leu Met Arg Gly Leu Lys Pro Trp Thr Gln
545 550 555 560

Tyr Ala Ile Phe Val Lys Thr Leu Val Thr Phe Ser Asp Glu Arg Arg
565 570 575

Thr Tyr Gly Ala Lys Ser Asp Ile Ile Tyr Val Gln Thr Asp Ala Thr
580 585 590

Asn Pro Ser Val Pro Leu Asp Pro Ile Ser Val Ser Asn Ser Ser Ser
595 600 605

Gln Ile Ile Leu Lys Trp Lys Pro Pro Ser Asp Pro Asn Gly Asn Ile
610 615 620

Thr His Tyr Leu Val Phe Trp Glu Arg Gln Ala Glu Asp Ser Glu Leu
625 630 635 640

Phe Glu Leu Asp Tyr Cys Leu Lys Gly Leu Lys Leu Pro Ser Arg Thr
645 650 655

Trp Ser Pro Pro Phe Glu Ser Glu Asp Ser Gln Lys His Asn Gln Ser
660 665 670

Glu Tyr Glu Asp Ser Ala Gly Glu Cys Cys Ser Cys Pro Lys Thr Asp
675 680 685

Ser Gln Ile Leu Lys Glu Leu Glu Glu Ser Ser Phe Arg Lys Thr Phe
690 695 700

Glu Asp Tyr Leu His Asn Val Val Phe Val Pro Arg Pro Ser Arg Lys
705 710 715 720

Arg Arg Ser Leu Gly Asp Val Gly Asn Val Thr Val Ala Val Pro Thr
725 730 735

Val Ala Ala Phe Pro Asn Thr Ser Ser Thr Ser Val Pro Thr Ser Pro
740 745 750

Glu Glu His Arg Pro Phe Glu Lys Val Val Asn Lys Glu Ser Leu Val
755 760 765

Ile Ser Gly Leu Arg His Phe Thr Gly Tyr Arg Ile Glu Leu Gln Ala
770 775 780

Cys Asn Gln Asp Thr Pro Glu Glu Arg Cys Ser Val Ala Ala Tyr Val

785					790					795					800
Ser	Ala	Arg	Thr	Met	Pro	Glu	Ala	Lys	Ala	Asp	Asp	Ile	Val	Gly	Pro
				805					810					815	
Val	Thr	His	Glu	Ile	Phe	Glu	Asn	Asn	Val	Val	His	Leu	Met	Trp	Gln
			820					825					830		
Glu	Pro	Lys	Glu	Pro	Asn	Gly	Leu	Ile	Val	Leu	Tyr	Glu	Val	Ser	Tyr
		835					840					845			
Arg	Arg	Tyr	Gly	Asp	Glu	Glu	Leu	His	Leu	Cys	Val	Ser	Arg	Lys	His
		850					855					860			
Phe	Ala	Leu	Glu	Arg	Gly	Cys	Arg	Leu	Arg	Gly	Leu	Ser	Pro	Gly	Asn
865						870				875					880
Tyr	Ser	Val	Arg	Ile	Arg	Ala	Thr	Ser	Leu	Ala	Gly	Asn	Gly	Ser	Trp
				885					890					895	
Thr	Glu	Pro	Thr	Tyr	Phe	Tyr	Val	Thr	Asp	Tyr	Leu	Asp	Val	Pro	Ser
			900					905					910		
Asn	Ile	Ala	Lys												
			915												
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Arg	Gln	Leu	Glu	Asn	Cys	Ser	Val	Val	Glu	Gly	His	Leu	Gln	Ile	Leu
			20					25					30		
Leu	Met	Phe	Thr	Ala	Thr	Gly	Glu	Asp	Phe	Arg	Gly	Leu	Ser	Phe	Pro
		35					40					45			
Arg	Leu	Thr	Gln	Val	Thr	Asp	Tyr	Leu	Leu	Leu	Phe	Arg	Val	Tyr	Gly
	50					55					60				

Leu Glu Ser Leu Arg Asp Leu Phe Pro Asn Leu Ala Val Ile Arg Gly
65 70 75 80

Thr Arg Leu Phe Leu Gly Tyr Ala Leu Val Ile Phe Glu Met Pro His
85 90 95

Leu Arg Asp Val Ala Leu Pro Ala Leu Gly Ala Val Leu Arg Gly Ala
100 105 110

Val Arg Val Glu Lys Asn Gln Glu Leu Cys His Leu Ser Thr Ile Asp
115 120 125

Trp Gly Leu Leu Gln Pro Ala Pro Gly Ala Asn His Ile Val Gly Asn
130 135 140

Lys Leu Gly Glu Glu Cys Ala Asp Val Cys Pro Gly Val Leu Gly Ala
145 150 155 160

Ala Gly Glu Pro Cys Ala Lys Thr Thr Phe Ser Gly His Thr Asp Tyr
165 170 175

Arg Cys Trp Thr Ser Ser His Cys Gln Arg Val Cys Pro Cys Pro His
180 185 190

Gly Met Ala Cys Thr Ala Arg Gly Glu Cys Cys His Thr Glu Cys Leu
195 200 205

Gly Gly Cys Ser Gln Pro Glu Asp Pro Arg Ala Cys Val Ala Cys Arg
210 215 220

His Leu Tyr Phe Gln Gly Ala Cys Leu Trp Ala Cys Pro Pro Gly Thr
225 230 235 240

Tyr Gln Tyr Glu Ser Trp Arg Cys Val Thr Ala Glu Arg Cys Ala Ser
245 250 255

Leu His Ser Val Pro Gly Arg Ala Ser Thr Phe Gly Ile His Gln Gly
260 265 270

Ser Cys Leu Ala Gln Cys Pro Ser Gly Phe Thr Arg Asn Ser Ser Ser
275 280 285

Ile Phe Cys His Lys Cys Glu Gly Leu Cys Pro Lys Glu Cys Lys Val
290 295 300

Gly Thr Lys Thr Ile Asp Ser Ile Gln Ala Ala Gln Asp Leu Val Gly
305 310 315 320

Cys Thr His Val Glu Gly Ser Leu Ile Leu Asn Leu Arg Gln Gly Tyr
325 330 335

Asn Leu Glu Pro Gln Leu Gln His Ser Leu Gly Leu Val Glu Thr Ile
340 345 350

Thr Gly Phe Leu Lys Ile Lys His Ser Phe Ala Leu Val Ser Leu Gly
355 360 365

Phe Phe Lys Asn Leu Lys Leu Ile Arg Gly Asp Ala Met Val Asp Gly
370 375 380

Asn Tyr Thr Leu Tyr Val Leu Asp Asn Gln Asn Leu Gln Gln Leu Gly
385 390 395 400

Ser Trp Val Ala Ala Gly Leu Thr Ile Pro Val Gly Lys Ile Tyr Phe
405 410 415

Ala Phe Asn Pro Arg Leu Cys Leu Glu His Ile Tyr Arg Leu Glu Glu
420 425 430

Val Thr Gly Thr Arg Gly Arg Gln Asn Lys Ala Glu Ile Asn Pro Arg
435 440 445

Thr Asn Gly Asp Arg Ala Ala Cys Gln Thr Arg Thr Leu Arg Phe Val
450 455 460

Ser Asn Val Thr Glu Ala Asp Arg Ile Leu Leu Arg Trp Glu Arg Tyr
465 470 475 480

Glu Pro Leu Glu Ala Arg Asp Leu Leu Ser Phe Ile Val Tyr Tyr Lys
485 490 495

Glu Ser Pro Phe Gln Asn Ala Thr Glu His Val Gly Pro Asp Ala Cys
500 505 510

Gly Thr Gln Ser Trp Asn Leu Leu Asp Val Glu Leu Pro Leu Ser Arg

515 520 525

Thr Gln Glu Pro Gly Val Thr Leu Ala Ser Leu Lys Pro Trp Thr Gln
530 535 540

Tyr Ala Val Phe Val Arg Ala Ile Thr Leu Thr Thr Glu Glu Asp Ser
545 550 555 560

Pro His Gln Gly Ala Gln Ser Pro Ile Val Tyr Leu Arg Thr Leu Pro
565 570 575

Ala Ala Pro Thr Val Pro Gln Asp Val Ile Ser Thr Ser Asn Ser Ser
580 585 590

Ser His Leu Leu Val Arg Trp Lys Pro Pro Thr Gln Arg Asn Gly Asn
595 600 605

Leu Thr Tyr Tyr Leu Val Leu Trp Gln Arg Leu Ala Glu Asp Gly Asp
610 615 620

Leu Tyr Leu Asn Asp Tyr Cys His Arg Gly Leu Arg Leu Pro Thr Ser
625 630 635 640

Asn Asn Asp Pro Arg Phe Asp Gly Glu Asp Gly Asp Pro Glu Ala Glu
645 650 655

Met Glu Ser Asp Cys Cys Pro Cys Gln His Pro Pro Pro Gly Gln Val
660 665 670

Leu Pro Pro Leu Glu Ala Gln Glu Ala Ser Phe Gln Lys Lys Phe Glu
675 680 685

Asn Phe Leu His Asn Ala Ile Thr Ile Pro Ile Ser Pro Trp Lys Val
690 695 700

Thr Ser Ile Asn Lys Ser Pro Gln Arg Asp Ser Gly Arg His Arg Arg
705 710 715 720

Ala Ala Gly Pro Leu Arg Leu Gly Gly Asn Ser Ser Asp Phe Glu Ile
725 730 735

Gln Glu Asp Lys Val Pro Arg Glu Arg Ala Val Leu Ser Gly Leu Arg
740 745 750

His Phe Thr Glu Tyr Arg Ile Asp Ile His Ala Cys Asn His Ala Ala
755 760 765

His Thr Val Gly Cys Ser Ala Ala Thr Phe Val Phe Ala Arg Thr Met
770 775 780

Pro His Arg Glu Ala Asp Gly Ile Pro Gly Lys Val Ala Trp Glu Ala
785 790 795 800

Ser Ser Lys Asn Ser Val Leu Leu Arg Trp Leu Glu Pro Pro Asp Pro
805 810 815

Asn Gly Leu Ile Leu Lys Tyr Glu Ile Lys Tyr Arg Arg Leu Gly Glu
820 825 830

Glu Ala Thr Val Leu Cys Val Ser Arg Leu Arg Tyr Ala Lys Phe Gly
835 840 845

Gly Val His Leu Ala Leu Leu Pro Pro Gly Asn Tyr Ser Ala Arg Val
850 855 860

Arg Ala Thr Ser Leu Ala Gly Asn Gly Ser Trp Thr Asp Ser Val Ala
865 870 875 880

Phe Tyr Ile Leu Gly Pro Glu Glu Glu Asp Ala Gly Gly Leu His
885 890 895

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<212> PRT

<213> Homo sapiens

<400> 15

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Leu Asn

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<211> 11

<212> PRT

<213> Homo sapiens

<400> 16

Glu Gln Lys Leu Ile Ser Glu Glu Asp Leu Asn
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